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## Where's the need? the use of specialist mental health services in adolescence and young adulthood

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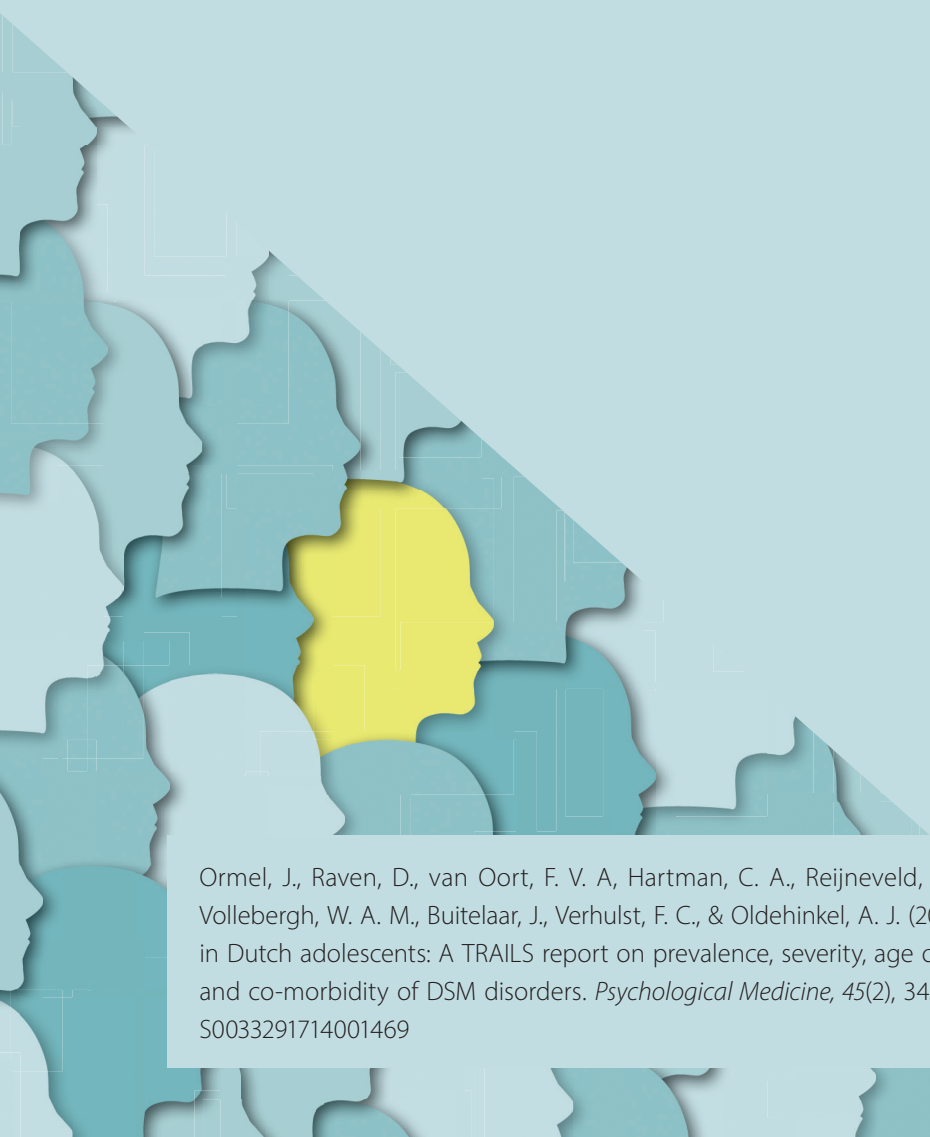
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# 2

## **Mental health in Dutch adolescents: A TRAILS report on prevalence, severity, age of onset, continuity and co-morbidity of DSM disorders**



Ormel, J., Raven, D., van Oort, F. V. A., Hartman, C. A., Reijneveld, S. A., Veenstra, R., Vollebergh, W. A. M., Buitelaar, J., Verhulst, F. C., & Oldehinkel, A. J. (2015). Mental health in Dutch adolescents: A TRAILS report on prevalence, severity, age of onset, continuity and co-morbidity of DSM disorders. *Psychological Medicine*, 45(2), 345-360. doi: 10.1017/S0033291714001469

## Abstract

**Background.** With psychopathology rising during adolescence and evidence suggesting that adult mental health burden is often due to disorders beginning in youth, it is important to investigate the epidemiology of adolescent mental disorders.

**Method.** We analyzed data gathered at ages 11 (baseline) and 19 years from the population-based Dutch TRacking Adolescents' Individual Lives Survey (TRAILS) study. At baseline we administered the Achenbach measures (Child Behavior Checklist, Youth Self-Report) and at age 19 years the World Health Organization's Composite International Diagnostic Interview version 3.0 (CIDI 3.0) to 1584 youths.

**Results.** Lifetime, 12-month and 30-day prevalences of any CIDI-DSM-IV disorder were 45, 31 and 15%, respectively. Half were severe. Anxiety disorders were the most common but the least severe whereas mood and behavior disorders were less prevalent but more severe. Disorders persisted, mostly by recurrence in mood disorders and chronicity in anxiety disorders. Median onset age varied substantially across disorders. Having one disorder increased subjects' risk of developing another disorder. We found substantial homotypic and heterotypic continuity. Baseline problems predicted the development of diagnosable disorders in adolescence. Non-intact families and low maternal education predicted externalizing disorders. Most morbidity concentrated in 5–10% of the sample, experiencing 34–55% of all severe lifetime disorders.

**Conclusions.** At late adolescence, 22% of youths have experienced a severe episode and 23% only mild episodes. This psychopathology is rather persistent, mostly due to recurrence, showing both monotypic and heterotypic continuity, with family context affecting particularly externalizing disorders. High problem levels at age 11 years are modest precursors of incident adolescent disorders. The burden of mental illness concentrates in 5–10% of the adolescent population.

**Key words:** Age of onset; Anxiety; Behavior disorders; Co-morbidity; Depression; Psychopathology

## 2.1 Introduction

Psychopathology is on the rise during adolescence (Rutter 1995, 2005; Newman *et al.* 1996) and evidence suggests that the adult mental health burden (eds Murray & Lopez 1996; Ormel *et al.* 2008) may be largely due to disorders with precursors or onset in childhood and adolescence (Kim-Cohen *et al.* 2003; Copeland *et al.* 2009). Because developmental pathways are set in motion or become entrenched during adolescence, adolescent psychopathology may have long-term consequences (Ferdinand *et al.* 1995; Quinton *et al.* 1995; Rutter & Maughan 1997; Costello *et al.* 1999; Fergusson & Horwood 2001; Verboom *et al.* 2014). Hence, it is important to understand the epidemiology of mental disorders during adolescence.

Earlier studies have yielded important information on many aspects of the epidemiology of mental disorders in children and adolescents (e.g. Costello *et al.* 1996, 2005; Verhulst *et al.* 1997; Angold *et al.* 1998; Fergusson & Horwood 2001; Ford *et al.* 2003; Maughan *et al.* 2008; Merikangas *et al.* 2010; Moffitt *et al.* 2010; Kessler *et al.* 2012a, 2012b). However, some important aspects remain unaddressed or need replication. These include severity, age of onset, persistence and continuity, and concentration of morbidity. Severity is important because it is unclear to what extent the previously reported remarkably high lifetime and 12-month prevalence rates represent mild disorders (Costello *et al.* 1996; Copeland *et al.* 2011; Kessler *et al.* 2012b). Age of onset and continuity are important issues as well. With a few exceptions (Kessler *et al.* 2011), age-of-onset information has rarely been used to its fullest potential, that is, by modelling age of onset as outcome or time-dependent covariate in a survival framework. Such a framework is highly appropriate to estimate the association of sociodemographic variables with mental disorder, adjusted for earlier disorders, and to study homotypic and heterotypic continuity of psychopathology. Homotypic continuity, in general, refers to the continuity of similar behaviors over time. In this paper, we analyze homotypic and heterotypic continuity of psychopathology at the level of classes of disorders (e.g. mood disorders) and the two broad domains of internalizing and externalizing disorders. Thus, homotypic continuity refers to continuity within class or domain whereas heterotypic continuity refers to continuity of psychopathology between classes or domains. Finally, concentration of morbidity is important because studies in adult populations suggest that in particular multimorbidity ( $\geq 3$  lifetime disorders) is associated with high levels of disability and service use (Kessler *et al.* 1994; Jenkins *et al.* 1997; Andrews *et al.* 2001; Jacobi *et al.* 2004).

The purpose of this paper, therefore, is to provide comprehensive epidemiological data on adolescent mental disorders. We distinguish four classes of disorders: anxiety, mood, behavior and substance use disorders. The first two belong to the internalizing domain, the last two to the externalizing domain. We are especially interested in the ratio of mild



to severe cases, age of onset, persistence (recurrence and chronicity), homotypic and heterotypic continuity, and the concentration of morbidity, and will also present data on prevalence (lifetime, 12-month, 30-day) and baseline problem levels and sociodemographic predictors analyzed in a multivariate survival framework.

## 2.2 Method

### Sample and procedure

The TRacking Adolescents' Individual Lives Survey (TRAILS) is a prospective cohort study of Dutch adolescents using bi- or triennial measurements from age 11 years onward. Its aim is to chart and explain the development of mental health from preadolescence into adulthood. Previous publications have extensively described its design, methods, and response rates and bias (de Winter *et al.* 2005; Huisman *et al.* 2008; Nederhof *et al.* 2012; Ormel *et al.* 2012). Briefly, participants were selected from five municipalities in the North of the Netherlands, both urban and rural areas, including the three largest cities. Children born between 1 October 1989 and 30 September 1991 were eligible for inclusion, providing their schools were willing to participate and they met the study's inclusion criteria (de Winter *et al.* 2005). Over 90% of the schools, enrolling a total of 2935 eligible children, agreed to participate in the study. Through extended efforts, 76% of these children and their parents consented to participate (T1,  $n=2230$ , mean age=11.1 years,  $SD=0.6$  years, 50.8% girls). Response rates at follow-ups ranged from 96.4% (T2,  $n=2149$ , mean age 13.6 9 years,  $SD=0.5$  years, 51.0% girls) to 81.4% (T3,  $n=1816$ , mean age 16.3 years,  $SD=0.7$  years, 52.3% girls). Each assessment wave was approved by the Dutch Central Committee on Research Involving Human Subjects (CCMO; [www.ccmo.nl](http://www.ccmo.nl)).

The data we present here were collected in the first (T1, baseline) and fourth (T4) assessment wave of TRAILS, which ran from March 2001 to July 2002 and from October 2008 to September 2010, respectively. The response rate at T4 was 84.3% of the initial T1 sample ( $n=1881$ , mean age 19.1 years,  $SD=0.6$  years, 52.3% girls) (Nederhof *et al.* 2012; Ormel *et al.* 2012). Not all T4 participants agreed to have the full diagnostic interview, but 1584 adolescents provided complete diagnostic data [Composite International Diagnostic Interview (CIDI), mean age 19.3 years, range 18–20 years, 54.0% girls], representing 84.2% of the T4 sample and 71.0% of the original T1 baseline sample. Response rates were somewhat better than for most European studies (Wittchen *et al.* 1998; Alonso *et al.* 2004a; de Graaf *et al.* 2012). Non-response was somewhat higher in males and in adolescents of non-Western ethnicity, with divorced parents, low socio-economic status (SES), low intelligence quotient and academic achievement, poor physical health, and with behavior and substance use problems (Nederhof *et al.* 2012). Multiple logistic regression analyses showed that these effects were partially overlapping. Non-response showed little to no association with

urbanization, parental religiousness, being a single child, or the most recently available self-reports of anxiety and mood problems.

### Sample representativeness

The TRAILS sample was largely (84.3%) collected from the three provincial capitals in the northern part of the Netherlands. This does not include the metropolitan area of the Randstad (Amsterdam, Rotterdam, Den Haag and Utrecht), which is more ethnically diverse. Apart from ethnicity and under-representation of people from extremely urbanized areas and – to a small extent – males, the T4 CIDI TRAILS sample is representative of the Dutch population aged 18–20 years (Table 2.1).

**Table 2.1.** Representativeness of the TRAILS sample

	National registries	TRAILS	
		Unweighted	Weighted
	%	%	%
<b>Population distribution (women)<sup>a</sup></b>	<b>49.0</b>	<b>54.0</b>	<b>51.1</b>
Marital status (married) <sup>a, b</sup>	0.3	0.2	0.2
Ethnicity (non-western) <sup>a</sup>	15.8	7.6	7.9
Parental net income (low; <€16,000) <sup>c</sup>	17.2	15.8	17.8
Urbanization degree (≥1,500 residential addresses per square kilometer) <sup>c</sup>	40.4	36.5	36.5

TRAILS, TRacking Adolescents' Individual Lives Survey.

<sup>a</sup> Census data and TRAILS sample data from 2009.

<sup>b</sup> Census data from ages 18–19.

<sup>c</sup> Census data and TRAILS sample data from 2001.

### Measures

#### *Diagnostic assessment*

TRAILS assessed the presence of mental disorders at T4 using the computer-assisted World Health Organization CIDI 3.0. The assessment included mood disorders (major depressive disorder, dysthymic disorder, and bipolar disorder I and II), anxiety disorders (panic disorder, agoraphobia, social phobia, specific phobia, generalized anxiety disorder, separation anxiety disorder, and obsessive–compulsive disorder), behavior disorders (attention-deficit/hyperactivity disorder, oppositional defiant disorder, and conduct disorder) and substance use disorders (alcohol abuse/dependence, drug abuse/dependence). TRAILS assessed eating disorders (anorexia nervosa, bulimia nervosa, binge-eating behavior) differently, so we have not included them.

The CIDI 3.0 is a structured diagnostic interview that has been used in multiple surveys worldwide to generate diagnoses based on the Diagnostic and Statistical Manual of Mental

Disorders, fourth edition (DSM-IV) (Kessler & Üstün 2004). The CIDI 3.0 assesses age of onset of any disorder with a series of questions that have been shown to yield plausible age-of-onset data (Kessler *et al.* 2005a). An important feature of the 3.0 version of the age-of-onset questions is the help of mnemonic aids and the sequence of onset questions, typically starting with the worst episode ever of the index disorder (when did it occur), followed by the most recent episode (when did it occur), and finally targeting the first ever episode and its age of onset (Kessler *et al.* 2005a).

In TRAILS, trained lay interviewers performed the CIDI at T4. Some clinical calibration studies found the CIDI's assessment of the selected disorders to be generally valid in comparison with blinded clinical reappraisal interviews using the Structured Clinical Interview for DSM-IV (SCID) (Kessler & Üstün 2004; Haro *et al.* 2006; Kessler *et al.* 2009) but in comparison with the Schedule for Clinical Assessment in Neuropsychiatry (SCAN) the CIDI performed less well (Brugha *et al.* 2001). CIDI-based prevalence estimates were typically no higher than SCID estimates, except for specific phobias and oppositional defiant disorder, but higher than SCAN estimates. The definitions of all disorders in the Dutch CIDI adhered to DSM-IV criteria. Diagnostic hierarchy rules were applied for every disorder, with the exception of substance use disorders. Impairment criteria embedded in the CIDI-DSM-IV diagnostic thresholds require the presence of at least some impairment or moderate symptom severity (distress) to make a diagnosis.

### ***Prevalence rates and ratios***

We established lifetime, 12-month and 30-day prevalence rates according to the DSM-IV (American Psychiatric Association 1994). In addition, we calculated the ratio of the 12-month prevalence to the lifetime prevalence, as well as the ratio of the 30-day prevalence to the 12-month prevalence. The ratio of 12-month prevalence to lifetime prevalence of a particular disorder tells – with certain assumptions on age of onset – something about its persistence. The 30-day to 12-month prevalence ratio tells something about the source of persistence: when smaller than the 12-month to lifetime prevalence ratio, it points at recurrence; when larger it points at chronicity.

### ***Severe disorders***

To separate mild from severe disorders, we used the Merikangas *et al.* (2010) definition of severe disorders. This definition sets higher thresholds for impairment and symptom severity than the CIDI-DSM-IV. To be severe, anxiety or mood disorders required both severe distress and impairment of daily activities. We did not separate agoraphobia and panic disorder into severe and less severe disorders because, following Merikangas *et al.* (2010), we considered the standard CIDI-DSM-IV severity rating for these disorders to be sufficiently severe. Behavior disorders required severe impairment to be classified as

severe. With regard to substance use disorders, we considered dependence severe and abuse non-severe unless it developed into dependence. The reason for this is that CIDI-DSM-IV substance use disorder in Dutch young people rarely is associated with functional impairment or distress (Bijl & Ravelli 2000; ten Have *et al.* 2013b).

### ***Baseline psychopathology***

The parent-report Child Behavior Checklist (CBCL) and the self-report Youth Self-Report (YSR) are questionnaires of good reliability and validity (Verhulst *et al.* 1997; Achenbach & Rescorla 2006) that cover behavioral and emotional problems in the past 6 months. Both contain about 112 problem items, which are scored on a three-point scale. Both consist of eight narrowband scales. In order to improve the match with DSM-IV diagnoses, Achenbach *et al.* (2003) constructed CBCL/YSR/DSM-IV scales. As a result, six CBCL/YSR/DSM-IV scales were derived: affective problems, anxiety problems, somatic problems, attention deficit/hyperactivity problems, oppositional defiant problems and conduct problems. These were used in the present study. Scale scores were dichotomized [normal range versus (sub)clinical range].

### ***Sociodemographic variables***

We measured the following sociodemographic variables at baseline: gender; age; ethnicity (Western origin, non-Western origin); SES, a composite measure of paternal and maternal education (elementary education, lower tracks of secondary education, higher tracks of secondary education, senior vocational training, university), occupation and family income (lowest 25%, middle 50%, highest 25%) (Veenstra *et al.* 2006); urbanicity [0–999 addresses per km<sup>2</sup> (low), 1000–2499 addresses per km<sup>2</sup> (moderate/strong), 2500 or more addresses per km<sup>2</sup> (extreme)] (Reijneveld *et al.* 2010); number of biological parents living with the respondent (both, not both); siblings (no, yes); and parental religiosity (non-religious, passively religious, actively religious) (van der Jagt-Jelsma *et al.* 2011).

### ***Statistical analysis***

To obtain weighted prevalence rates (Table 2.2), we used a sampling weight based on three indicators from the first measurement wave: gender, SES, and total problems score on the CBCL (normal, subclinical, clinical) to adjust for selective attrition (Achenbach & Rescorla 2006). The sample weight of cases with missing CBCL or SES information ( $n=95$ ; 6.0%) was set to 1. With the age-of-onset data, we generated standardized cumulative prevalence curves (Figure 2.1). Homotypic continuity, especially persistence of a disorder and whether it was due to recurrence or chronicity, was examined using prevalence ratios (Table 2.2). We used a multivariate Cox proportional hazards model (1) to analyze heterotypic continuity by (a) adding the onset of co-morbid disorders as time-dependent covariates (Table 2.3)

and (b) by linking baseline (age 11 years) problem levels to the onset of post-baseline disorders, and (2) to examine sociodemographic predictors (Table 2.4). Thus, effects of a particular predictor were adjusted for other predictors (e.g. other disorders in Table 2.3; and other sociodemographic covariates in Table 2.4). Our study evaluated all tests at the 0.05 significance level with two-sided tests.

## 2.3 Results

### Prevalence

Table 2.2 presents prevalence rates for CIDI-DSM-IV mental disorders by time-frame (lifetime, 12-month, 30-day) and severity. All four DSM classes of disorders were important components of overall lifetime prevalence. According to the lifetime time-frame, mood disorders affected 17% of the total sample: 15% met criteria for major depression. About one in four adolescents met criteria for an anxiety disorder, with rates for individual disorders ranging from 1% for agoraphobia without panic disorder to 12% for specific and social phobia. Behavior disorders affected 16% of the sample, with about equal rates for oppositional defiant and conduct disorder. Prevalence rates for substance dependence were substantially lower than for substance abuse. Nearly 45% of the total sample experienced at least one of the disorders in Table 2.2 during their lives, with 5.2% of the sample having disorders from 53 different classes and 10.1% of the sample having three or more disorders lifetime irrespective of class.

### Severe disorders

The lifetime prevalence of severe disorders was 22%; for half of the total lifetime prevalence, 23% were mild. In general, mood and behavior disorders were more often severe than anxiety disorders (Table 2.2). Severe mood disorders represented 49% of all mood disorders, while severe anxiety disorders represented only 19% of all anxiety disorders. Severe anxiety cases included relatively many individuals with generalized anxiety, obsessive–compulsive disorder, panic disorder and agoraphobia. Cases of separation anxiety disorder, specific phobia and social phobia were typically milder. Severe behavior disorders comprised nearly a third of all the severe cases in the sample. The proportion of subjects with at least one severe disorder rose with increasing co-morbidity across classes, from 29% for respondents with only one disorder to 96% for respondents with disorders from 53 different classes.

**Table 2.2.** Weighted<sup>a</sup> prevalences, prevalence ratios and age of onset of DSM-IV disorders in TRAILS (*n*=1,584)

	Prevalence				Ratio Severe / Lifetime	Prevalence ratios		Age-of-onset		
	30 Days	12 Months	Lifetime	Severe Lifetime		12 Month / Lifetime	30 Days / 12 Month	Mean (S.E.)	Median	IQR
	% (S.E.)	% (S.E.)	% (S.E.)	% (S.E.)						
Mood disorders										
Bipolar I disorder	0.2 (0.1)	0.2 (0.1)	0.4 (0.2)	0.1 (0.1)	25.6	51.3	74.7	14.6 (1.0)	15	5
Bipolar II disorder	0.5 (0.2)	0.9 (0.2)	1.1 (0.3)	0.6 (0.2)	53.7	83.2	55.2	15.1 (0.8)	16	2
Major depressive disorder	2.2 (0.4)	8.8 (0.7)	15.5 (0.9)	7.5 (0.7)	48.2	56.6	25.0	14.1 (0.2)	14	4
Dysthymia	0.5 (0.2)	1.6 (0.3)	1.7 (0.3)	1.1 (0.3)	63.1	93.4	31.0	13.9 (0.6)	14	4
Any mood disorder	2.9 (0.4)	10.2 (0.8)	17.3 (1.0)	8.4 (0.7)	48.5	58.8	28.7	14.2 (0.2)	15	4
Anxiety disorders										
Separation anxiety disorder	0.1 (0.1)	0.3 (0.1)	3.1 (0.4)	0.3 (0.1)	11.2	9.8	40.3	9.1 (0.6)	7	9
Agoraphobia (without PAN)	0.1 (0.1)	0.7 (0.2)	1.0 (0.2)	1.0 (0.2) <sup>b</sup>	100.0	73.6	17.1	11.6 (1.1)	12	8
Generalized anxiety disorder	0.7 (0.2)	1.8 (0.3)	2.9 (0.4)	0.9 (0.2)	31.2	62.2	36.8	13.2 (0.5)	14	4
Obsessive compulsive disorder	2.2 (0.4)	3.4 (0.5)	5.9 (0.6)	0.9 (0.2)	15.5	56.9	66.5	11.5 (0.5)	13	9
Panic disorder	0.3 (0.1)	1.3 (0.3)	1.6 (0.3)	1.6 (0.3) <sup>b</sup>	100.0	79.1	24.3	13.7 (0.8)	15	6
Social phobia	3.2 (0.4)	7.5 (0.7)	12.4 (0.8)	0.9 (0.2)	7.3	60.2	42.8	10.1 (0.3)	11	5
Specific phobia	5.6 (0.6)	9.0 (0.7)	11.5 (0.8)	0.5 (0.2)	4.7	78.0	62.2	6.8 (0.3)	5	4
Any anxiety disorder	10.6 (0.8)	18.4 (1.0)	28.0 (1.1)	5.2 (0.6)	18.7	65.8	57.8	8.8 (0.2)	8	8
Behavior disorders										
Attention deficit disorder	–	3.2 (0.4)	4.2 (0.5) <sup>c</sup>	1.6 (0.3)	37.7	76.4	– <sup>c</sup>	5.4 (0.2)	5	2
Oppositional defiant disorder	–	1.4 (0.3)	8.9 (0.7) <sup>c</sup>	4.7 (0.5)	53.4	16.2	– <sup>c</sup>	10.2 (0.3)	11	6
Conduct disorder	–	4.2 (0.5)	8.6 (0.7) <sup>c</sup>	4.3 (0.5)	49.5	48.7	– <sup>c</sup>	11.0 (0.3)	12	6
Any behavior disorder	–	7.6 (0.7)	16.2 (0.9) <sup>c</sup>	8.4 (0.7)	51.5	47.1	– <sup>c</sup>	9.0 (0.2)	8	8
Substance disorders										
Alcohol abuse	8.3 (0.7)	18.4 (1.0)	25.1 (1.1)	2.6 (0.4) <sup>d</sup>	10.5	73.2	45.4	16.1 (0.1)	16	2
Drug abuse	2.7 (0.4)	6.8 (0.6)	13.2 (0.9)	4.2 (0.5) <sup>d</sup>	32.2	51.6	39.8	16.0 (0.1)	16	2

**Table 2.2 (Continued).** Weighted<sup>a</sup> prevalences, prevalence ratios and age of onset of DSM-IV disorders in TRAILS (n=1,584)

	Prevalence			Ratio Severe / Lifetime	Prevalence ratios		Age-of-onset	
	30 Days	12 Months	Lifetime		12 Month / Lifetime	30 Days / 12 Month	Mean (S.E.)	Median
	% (S.E.)	% (S.E.)	% (S.E.)					IQR
Any substance abuse	10.3 (0.8)	21.6 (1.0)	29.9 (1.2)	6.3 (0.6) <sup>d</sup>	21.0	72.3	15.9 (0.1)	16
Alcohol dependence	1.3 (0.3)	2.5 (0.4)	3.2 (0.4)	3.2 (0.4) <sup>b</sup>	100.0	80.3	16.8 (0.2)	17
Drug dependence	1.1 (0.3)	2.7 (0.4)	4.5 (0.5)	4.5 (0.5) <sup>b</sup>	100.0	59.0	16.3 (0.2)	16
Any substance dependence	2.3 (0.4)	4.9 (0.5)	7.1 (0.6)	7.1 (0.6) <sup>b</sup>	100.0	69.2	16.5 (0.1)	17
<b>Total classes (excl substance abuse)</b>								
Any class	14.5 (0.9)	31.0 (1.2)	44.8 (1.2)	21.9 (1.0)	49.0	69.3	9.5 (0.2)	9
Exactly 1 class	13.3 (0.9)	22.9 (1.1)	27.2 (1.1)	7.9 (0.7)	29.0	84.0	10.4 (0.2)	11
Exactly 2 classes	1.1 (0.3)	6.4 (0.6)	12.4 (0.8)	9.1 (0.7)	73.5	52.1	8.8 (0.3)	8
3 or 4 classes	0.1 (0.1)	1.7 (0.3)	5.2 (0.6)	5.0 (0.5)	95.8	33.0	6.8 (0.4)	6

S.E. = standard error; PAN = panic disorder; IQR = interquartile range.

<sup>a</sup> Cases weighted by gender, Child Behavior Check List (CBCL) cut-offs (normal, borderline clinical and clinical) and parental socioeconomic status (SES). Cases with missing CBCL and/or SES were assigned the weight 1.

<sup>b</sup> All lifetime disorders meet criteria for severe lifetime disorder.

<sup>c</sup> 30-days prevalence not established.

<sup>d</sup> Severe substance abuse defined as substance dependence.

### Age of onset

Figure 2.1 shows the standardized cumulative prevalence graphs. Major depressive disorder, dysthymia, and bipolar I and II are combined, and so are the phobias, and the other anxiety disorders except separation anxiety. The curves track the lifetime prevalence of each index disorder at each age. We standardized each curve as a proportion of its lifetime prevalence at age 19 years, which reduced between-disorder variations in prevalence to ease comparisons between ages of onset. The curves of disorders of the same class are the same colour. Visual approximation of these data distinguishes seven age-of-onset groups. These onset groups, which do not overlap with the four classes of disorder, are as follows:

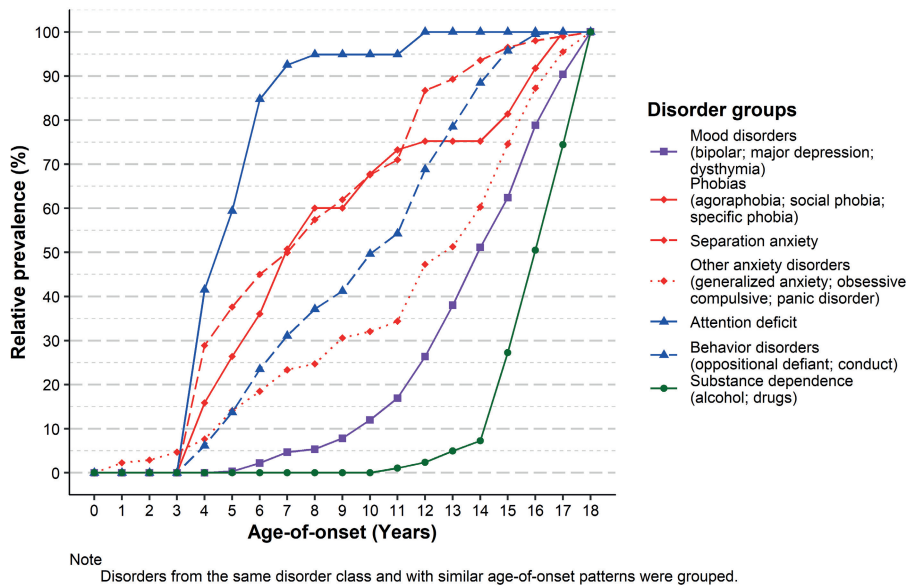
- 1) Attention deficit/hyperactivity disorder occurred earliest; onsets increase rapidly in early childhood, with virtually no new onset after age 6 years.
- 2) Phobia had early onsets as well. Most phobias, especially the specific phobias, had onsets before age 8 years and virtually no new onset occurred after age 14 years.
- 3) Separation anxiety closely followed phobia with one difference: new onsets occurred until age 17 years except during age 11–14 years when hardly any onset of separation anxiety occurred.
- 4) Behavior disorders began around the time of school entry and their onsets increased steadily until age 14–15 years.
- 5) Other anxiety disorders (generalized anxiety disorder, obsessive–compulsive disorder, panic disorder) tended to develop on average 2 years later than the behavior disorders; they were not prevalent until early adolescence, after which their incidence rose steadily.
- 6) Mood disorders were even less prevalent until early adolescence, after which their incidence rose steadily as well. Bipolar disorder had a slightly later onset.
- 7) Drug and alcohol dependence had the latest age of onset, with incidences beginning at age 14 years and steadily increasing after that.

Table 2.2 shows the mean and median age of onset for each disorder.

### Homotypic continuity

As shown in Table 2.2, the overall 12-month prevalence was 31%, which represented 69% of lifetime prevalence, while the 30-day prevalence was 14%, 47% of the 12-month prevalence. The ratio of 12-month prevalence to lifetime prevalence showed a wide range across disorders: from 10% for separation anxiety to 93% for dysthymia. The interquartile range was 52–76%, suggesting substantial persistence. The 30-day to 12-month prevalence ratios were typically smaller than the 12-month to lifetime prevalence ratios with only a few exceptions, suggesting that, on the whole, within-class continuity (persistence) comes more from recurrence than chronicity.





**Figure 2.1.** Standardized cumulative prevalence curves for Diagnostic and Statistical Manual of Mental Disorders, fourth edition (DSM-IV) disorders

## Heterotypic continuity

As expected, the presence of a mental disorder substantially increased the subject's risk of developing a disorder of a different class (Table 2.3). Of the 12 hazard ratios tested, 11 were significant, ranging from 2 to 5. The exception was anxiety disorders, which did not increase the risk of substance dependence. We found the strongest heterotypic continuity, in both directions, between behavior disorders and substance dependence.

## Baseline problems predict onset of disorders in adolescence

The previous continuity analyses were all based on retrospectively collected CIDI data. To supplement this with prospective data, we examined the predictive value of (sub) clinical baseline emotional and behavior problems as assessed at age 11 years with CBCL (parent-report) and the YSR (self-report) with regard to the post-baseline onset of CIDI-DSM-IV disorders (Appendix Tables A2.1 and A2.2). Because all attention deficit disorders, most specific phobia and separation anxiety disorders, and many oppositional disorders had an onset prior to baseline, they are not included in the post-baseline onset group. To compensate for this, we also linked baseline problems to the 12-month prevalence at age 19 years (Appendix Tables A2.3 and A2.4). We found substantial continuity at the level of the broad domains of internalizing and externalizing problems; at the disorder-class

level, continuity was less marked. Mood and anxiety disorders were predicted by baseline affective and anxiety problems; behavior disorders by baseline oppositional, conduct and affective problems whereas baseline anxiety problems reduced the risk of behavior disorders. Substance dependence was predicted by conduct, affective and attention problems. Effects were typically weak with most hazard ratios in the 1.5–2.5 range but it should be noted that effects of all baseline problem scales were adjusted for each other. We obtained similar results for the 12-month prevalence at age 19 years, with the self-report YSR being a better predictor than the parent-reported CBCL. The latter showed only a few significant associations with the 12-month prevalence of disorders, with the association between (sub)clinical baseline attention problems and any behavior disorder being the strongest (odds ratio 3.83, 95% confidence interval 2.17–6.75).

### **Sociodemographic predictors**

Table 2.4 presents the adjusted hazard ratios of the selected sociodemographic characteristics assessed at baseline for each class of mental disorder. We found the most significant associations between sociodemographic variables and behavior disorders. Associations of sociodemographic variables with mood, anxiety and substance use disorders were typically non-significant or weak. The strongest associations were found for gender, SES, and absence of one or both biological parents. Men had a substantially lower risk for anxiety and mood disorders than women, but a significantly higher risk of behavior disorders. The smaller than unity gender $\times$ time interaction indicates that the effect of gender on risk for behavior disorders decreased during adolescence while the larger than unity ethnicity $\times$ time interaction indicates that the effect of ethnicity increases. Maternal education accounted for most of the SES effect on behavior disorders. Neither parental income nor professional status, the other components of SES, predicted much change in mental health risks (data available on request). Urbanization predicted only behavior disorders which were more prevalent in highly urbanized areas.

### **Concentration of morbidity**

Nearly 75% of lifetime disorders were co-morbid disorders. Table 2.5 shows that the concentration of morbidity in adolescents with lifetime disorders from multiple classes is highly prominent. The 5.2% of the sample with a lifetime history of disorders from 53 classes accounts for a third of all severe lifetime disorders and slightly more than a quarter of all 12-month and 30-day disorders. Concentration of morbidity was relatively similar among the 10.1% with 53 disorders irrespective of class who accounted for 55% of all severe lifetime disorders and nearly half of all 12-month and 30-day disorders.

**Table 2.3.** HR estimates from Cox regression analyses of co-morbidity on age of onset of DSM-IV disorders by class (n=1,558)<sup>a</sup>

	Any mood disorder		Any anxiety disorder		Any behavior disorder		Any substance dependence	
	HR	95% CI	HR	95% CI	HR	95% CI	HR	95% CI
<b>DSM-IV disorders<sup>b</sup></b>								
Any mood disorder	–		2.66 ***	(1.82-3.89)	2.05 **	(1.19-3.53)	2.69 ***	(1.77-4.08)
Any anxiety disorder (without specific phobia)	2.97 ***	(2.30-3.83)	–		2.36 ***	(1.68-3.32)	0.88	(0.56-1.37)
Any behavioral disorder	2.07 ***	(1.55-2.75)	2.07 ***	(1.54-2.78)	–		4.90 ***	(3.32-7.23)
Any dependence disorder	1.67 ~	(0.91-3.08)	2.98 **	(1.50-5.90)	4.65 *	(1.10-19.58)	–	
<b>Model characteristics</b>								
Number of onsets	268		327		239		109	
Model improvement (chi-sq; df)	105.8 ***	3	59.2 ***	3	33.5 ***	3	94.9 ***	3

HR = Hazard ratio; CI = Confidence interval; chi-sq = Chi squared; df = Degrees of freedom.

<sup>a</sup> DSM-IV hierarchy rules applied where applicable.

<sup>b</sup> Aggregate DSM-IV any disorders added as time dependent covariates (ref = no onset before time T). Any disorders include the disorders as listed in Table 2.2 (excl specific phobia).

~ p<0.10 \* p<0.05 \*\* p<0.01 \*\*\* p<0.001

**Table 2.4.** HR estimates from multiple Cox regression analyses of T1 demographic covariates on age of onset of DSM-IV disorders by class ( $n=1,558$ )<sup>a,b</sup>

	Any mood disorder			Any anxiety disorder			Any behavior disorder			Any substance dependence		
	HR	95% CI	HR	95% CI	HR	95% CI	HR	95% CI	HR	95% CI	HR	95% CI
<b>Demographic covariates<sup>c,d</sup></b>												
Gender (ref = female)	0.49 ***	(0.38-0.63)	0.51 ***	(0.42-0.63)	2.13 ***	(1.59-2.84)	1.30	(0.87-1.92)				
Gender * Time	–		–		0.88 ***	(0.82-0.95)	–					
Ethnicity (ref = Dutch or other western country)	0.49	(0.16-1.49)	1.00	(0.70-1.43)	0.82	(0.46-1.47)	0.98	(0.48-2.00)				
Ethnicity * Time	1.16 *	(1.00-1.35)	–		1.14 *	(1.01-1.29)	–					
Socio-economic status (ref = high)												
Low socio-economic status	1.19	(0.83-1.70)	1.05	(0.79-1.38)	1.57 *	(1.09-2.25)	0.65	(0.39-1.09)				
Middle socio-economic status	1.04	(0.78-1.40)	1.05	(0.84-1.31)	1.29	(0.94-1.78)	0.61 *	(0.40-0.94)				
Parental religiosity (ref = not religious)												
At least one parent passive religious	1.17	(0.88-1.55)	1.03	(0.82-1.29)	0.96	(0.71-1.30)	1.09	(0.72-1.64)				
At least one parent active religious	0.98	(0.71-1.37)	0.92	(0.71-1.19)	0.80	(0.56-1.14)	0.24 ***	(0.10-0.56)				
Not both biological parents in the household (ref = both)	1.29 ~	(0.98-1.70)	1.15	(0.92-1.44)	1.53 **	(1.15-2.03)	1.37	(0.90-2.08)				
Single child (ref = has siblings)	0.94	(0.60-1.45)	1.27	(0.91-1.78)	0.80	(0.50-1.28)	0.98	(0.51-1.88)				
Urbanization (ref = low)												
Moderate urbanization	1.09	(0.82-1.44)	1.23 ~	(0.98-1.55)	1.01	(0.74-1.37)	1.06	(0.68-1.67)				
Extreme urbanization	0.73	(0.47-1.12)	0.96	(0.68-1.35)	1.53 *	(1.05-2.25)	1.12	(0.64-1.98)				
Moderate urbanization * Time	–		0.94 *	(0.90-0.99)	–		–					
Extreme urbanization * Time	–		1.00	(0.93-1.07)	–		–					
<b>Model characteristics</b>												
Number of onsets	268		432		239		109					
Model improvement (chi-sq; df)	159.6 ***	14	116.7 ***	15	98.5 ***	15	125.7 ***	13				

HR = hazard ratio; CI = confidence interval; ref = reference category; chi-sq = chi squared; df = degrees of freedom.

<sup>a</sup> DSM-IV hierarchy rules applied where applicable.<sup>b</sup> All models include aggregate DSM-IV disorders from other classes as time dependent covariates (ref = no onset before time T).<sup>c</sup> Demographic covariates measured at T1 (age 10-12).<sup>d</sup> Demographics entered as time independent covariates. If the proportional hazards assumption was violated interactions with time centered on age 11 (mean age at T1) were added.~  $p<0.10$  \*  $p<0.05$  \*\*  $p<0.01$  \*\*\*  $p<0.001$

**Table 2.5.** Clustering of lifetime, severe lifetime, 12-month and 30-day disorders among persons with lifetime co-morbidity

	Proportion of sample	Proportion of lifetime disorders	Proportion of severe lifetime disorders	Proportion of 12 month disorders	Proportion of 30 day disorders
	% (S.E.)	% (S.E.)	% (S.E.)	% (S.E.)	% (S.E.)
<b>Number of lifetime classes</b>					
0 classes	55.2 (1.2)	–	–	–	–
1 class	27.2 (1.1)	38.3 (1.3)	25.7 (1.9)	38.0 (1.7)	38.9 (2.9)
2 classes	12.4 (0.8)	35.2 (1.3)	40.5 (2.1)	35.1 (1.7)	35.1 (2.8)
3 or 4 classes	5.2 (0.6)	26.4 (1.2)	33.8 (2.0)	26.9 (1.6)	26.1 (2.6)

S.E. = standard error.

## 2.4 Discussion

### Strengths and limitations

Our findings should be interpreted in the light of strengths and limitations. Strengths of this study include its well-documented sample of adolescents, followed from preadolescence to adulthood and the considerable sample size. One limitation is that, despite limited non-response at baseline and attrition at follow-ups, CIDI non-response was significant and not entirely random. Bias due to non-response in psychiatric epidemiological studies tends to be conservative, with actual prevalence rates often higher and actual associations stronger, especially for externalizing disorders (Eaton *et al.* 1994; Kessler *et al.* 2005b; Merikangas *et al.* 2010a). Another limitation arises from the fact that CIDI-DSM-IV diagnoses were based on fully structured lay interviews carried out at age 19 years and not verified by professionals with clinical expertise. This very probably will have inflated prevalence estimates in comparison with semi-structured diagnostic interviews such as the SCAN and SCID (Brugha *et al.* 2001; Haro *et al.* 2006). On the other hand, TRAILS' single diagnostic CIDI assessment has probably resulted in deflated lifetime estimates, as studies with multiple cumulative diagnostic assessments report substantially higher prevalence rates than studies with a single CIDI administration (Moffitt *et al.* 2010; Copeland *et al.* 2011). Finally, we did not collect diagnostic information from sources other than the respondent, which might affect reliability (Ford *et al.* 2003). Evidence suggests, however, that this is not a major concern, as the reliability of self-reports increases during adolescence, while that of parents and teachers decreases (Edelbrock *et al.* 1985).

### Similar lifetime prevalence across Western countries

Direct and detailed comparisons with other adolescent studies are complicated by between-study differences in sampling, age range, and – very importantly – the DSM-IV

categories included. Our report did not include eating disorders, somatoform disorders or post-traumatic stress disorder. Nevertheless, after accounting for differences in included diagnoses, our findings are remarkably similar in overall prevalence rates to studies in industrialized countries that have used similar methodology (a single CIDI-DSM-IV assessment). Overall lifetime and 12-month prevalence in late adolescence tends to fluctuate around 45% and 30% (e.g. McGee *et al.* 1992; Wittchen *et al.* 1998; Merikangas *et al.* 2010), of which our findings suggest that about half are severe disorders.

### **Similar lifetime prevalence in youth and adults**

Most disorder-specific lifetime prevalence rates in TRAILS' adolescents approximate, and some even exceed, those found in nationally representative CIDI-DSM-IV surveys of adults (Kessler *et al.* 2005a; de Graaf *et al.* 2012). Two factors may explain this phenomenon. First, adults are more likely to forget earlier (mild) episodes or are unwilling to disclose them (Simon & VonKorff 1995; Moffitt *et al.* 2010). If this under-reporting increases and accumulates with age, lifetime prevalence may falsely appear to remain stable with increasing sample age. Second, retrospective evidence shows that many adult mental disorders, especially chronic–recurrent disorders, have early initial onsets during childhood and adolescence (Kessler *et al.* 1994, 2005a; Bijl *et al.* 1998), which would be detected by and included in prevalence surveys of adolescents.

### **Within-class homotypic continuity**

Most 12-month to lifetime prevalence ratios of individual disorders exceeded 0.60. Although confounding by recent onset and under-reporting of brief mild episodes is likely, these ratios suggest that most disorders are quite persistent. Consistent with earlier studies (Merikangas *et al.* 2010a; Kessler *et al.* 2012a), the 30-day to 12-month prevalence ratios were typically lower than 12-month to lifetime ratios, supporting the possibility that disorder persistence may be due more to episodic recurrence than to chronicity. Higher 30-day to 12-month ratios for anxiety disorders than for mood disorders suggest that anxiety disorders are more often chronic than mood disorders.

### **Between-class co-morbidity and heterotypic continuity**

Research on the structure of co-morbidity among common mental disorders has largely focused on prevalence (Angold *et al.* 1999), rather than on its development (Kessler *et al.* 2011). Using Cox regression analysis, we found moderate heterotypic continuity in both directions between all four classes of disorders except for anxiety to substance use. This similarity of heterotypic continuity between all disorder classes is interesting because one would expect that disorder classes that tend to onset early would be stronger predictors of classes tending to onset later, but not the other way around.

### **Baseline problem levels as domain-specific precursors**

The continuity findings were all based on retrospectively collected CIDI data. To supplement this we examined the predictive value of baseline problems for the onset of disorders in adolescence and the 12-month prevalence at age 19 years. We found domain-level homotypic continuity, i.e. baseline externalizing problems predicted later externalizing disorders and baseline internalizing problems predicted later internalizing disorders, and also heterotypic continuity as baseline internalizing problems predicted externalizing disorders (but not the other way around). Effects were weak to moderate, with a 1.5- to 3.05-fold increase in risk. Self-report of baseline problems was a better predictor of 12-month prevalence of disorders than parent-report, with the exception of parent-reported attention problems, which strongly predicted 12-month prevalence of behavior disorders. Collectively, these findings suggest that problem levels at age 11 years are weak to moderate predictors of the development of diagnosable disorders in adolescence.

### **Age of onset**

Although age-of-onset distributions varied between disorders, they definitely overlapped. New cases of each individual disorder, except for specific phobias and – by definition – attention deficit disorder, continued to develop throughout adolescence. Our age-of-onset findings confirm other reports (Kim-Cohen *et al.* 2003; Costello *et al.* 2005b; Merikangas *et al.* 2010a). These age-of-onset patterns are the opposite of those for nearly all chronic physical disorders, for which risks increase with age, peaking in late-middle and old age (van den Akker *et al.* 1998; Yach *et al.* 2004). Conversely, mental disorders tend to begin in youth, with substantially lower future risk for those who enter adulthood without any lifetime mental disorder (Kessler *et al.* 2005b).

### **Sociodemographic predictors**

Our results regarding sociodemographic variables are largely consistent with previous research (McGee *et al.* 1992; Costello *et al.* 1996, 2005a; Verhulst *et al.* 1997; Fergusson & Horwood 2001; Ford *et al.* 2003; Merikangas *et al.* 2010a; Kessler *et al.* 2012a). Gender was a strong correlate, with girls having more anxiety and mood disorders and fewer behavior disorders. Absence of one or both biological parents in the household and low SES, especially low maternal education, predicted behavior disorders, but not anxiety and mood disorders. Though the significance of parental education and family composition has already been well documented, including for childhood physical health outcomes (Merikangas *et al.* 2010a), the causal dynamics are still unclear (Fergusson & Horwood 2001; Shanahan *et al.* 2008).

## Clinical and public health implications

We observed substantial co-morbidity. Half of all affected youth had at least one additional lifetime diagnosis, 10% of the entire sample had three or more lifetime disorders, and 5% had lifetime disorders from three or four classes. These results strongly indicate that, even at this young age, co-morbidity between classes of disorders is not uncommon. Given the differences in included diagnoses, our co-morbidity rates are roughly similar to those reported for German and US adolescents and young adults (Wittchen *et al.* 1998; Kessler *et al.* 2012c). Co-morbidity was associated with high overall severity, as 10% of the sample with  $\geq 3$  disorders and the 5% of the sample with disorders from  $\geq 3$  classes experienced 55% and 34% of all severe lifetime disorders, respectively.

The observed concentration of morbidity is consistent with the possibility of a general psychopathology severity dimension. During the past two decades, strong evidence has shown three underlying dimensions to psychopathology, which represent the core psychopathological liabilities (or processes) of internalizing, externalizing and thought disturbance (Vollebergh *et al.* 2001; Kotov *et al.* 2011; Krueger & Markon 2011). Historically, the possibility of a superordinate general psychopathology severity dimension has been suggested (Wing *et al.* 1978). Empirical support for a general severity dimension for psychopathology continues to accumulate (Lahey *et al.* 2012; Caspi *et al.* 2014).

The concentration of morbidity has diagnostic implications as well (e.g. Kessler *et al.* 2012c; Uher & Rutter 2012). Is the splitting up of symptom clusters into many different disorders in the DSM-IV, and the DSM-5 as well, correct and helpful? Are those with severe co-morbidity the unlucky few who have developed multiple separate disorders or are we struggling with a syndrome uncharacterized by current classifications? From a public health and aetiological perspective, splitting into specific disorders may be less effective than trans-diagnostic classification at the disorder-class level, especially if age-of-onset patterns are taken into account. From a clinical perspective, the answer depends strongly on the value to treatment of distinguishing between specific disorders.

The high prevalence of mild CIDI-DSM-IV disorders raises two related questions: does the CIDI-DSM-IV overdiagnose – and – given that mild disorders in adolescence predict serious adult disorders, do mild cases require intervention (Kessler *et al.* 2003)? If treatment of mild disorders in childhood reduces future risk – an untested assumption as yet – then mild disorders should be treated and attempts to avoid overdiagnosis are actually unwanted and the current diagnostic cutoff between normal variation and mild disorder has utility. If, however, treatment of mild disorders does not reduce future risk, then the resources are better spent on prevention and treatment of severe disorders and co-morbidity. In this case, the current diagnostic cut-off between normal variation and mild disorder has less utility.



The fact that the burden of psychopathology is concentrated in youths with multiple lifetime disorders suggests focusing treatment and prevention on youth with lifetime multimorbidity. Unfortunately, because clinical trials in children and adolescents are relatively rare and in addition tend to exclude co-morbid cases, the evidence on prevention and treatment of multimorbidity is virtually lacking. To improve long-term outcomes, early prevention and treatment programmes perhaps best target self-control and neuroticism in addition to the mental disorders as these temperamental traits seem to play an important role in the development of co-morbidity and associated life outcomes (Oldehinkel *et al.* 2004; Lahey 2009; Moffitt *et al.* 2011; Ormel *et al.* 2013).

## **Conclusions**

Our findings, supported by earlier evidence, justify four conclusions about the mental health of adolescents in Western populations. First, as shown by prospective cumulative studies (Moffitt *et al.* 2010; Copeland *et al.* 2011), episodes of mild DSM-IV mental disorder are common. In that respect, mental illness is no different from physical illness, with its common episodes of influenza, colds, migraine and injuries. The second conclusion stresses that in slightly over half of the lifetime DSM-IV disorder prevalence, the disorder is mild, but the third conclusion emphasizes that a fifth of the adolescents experienced at least one severe disorder. Notably, the prevalence of severe mental disorder in adolescents is higher than even the most prevalent major somatic conditions, including asthma and diabetes (Eder *et al.* 2006; Hossain *et al.* 2007). The fourth conclusion highlights that about 10% of all youth have poor mental health and may be at risk of long-term mental illness in adulthood. Collectively, the findings point strongly to the need to investigate the long-term effects on adult mental health risk of (i) early and intensive treatment of multimorbid youth and (ii) non-intensive treatment of mild disorders.



